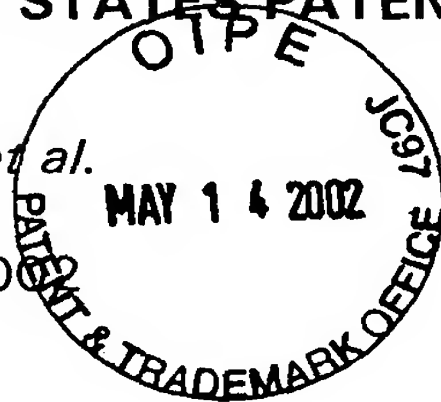


IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: FREDEKING *et al.*  
Serial No.: 10/038,557  
Filed: January 3, 2002  
Confirmation No.: 8399



For: *COMPOSITIONS AND METHODS  
FOR TREATING HEMORRHAGIC  
VIRUS INFECTIONS AND OTHER  
DISORDERS*

Art Unit: 1646  
Examiner: Unassigned

I hereby certify that this paper and the attached papers are being deposited with the United States Postal Service as first class mail in an envelope addressed to:

U.S. Patent and Trademark Office  
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Maritza O'Neill

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TRANSMITTAL LETTER

Commissioner for Patents  
Arlington, VA 22202

Dear Sir:

Transmitted herewith is a Preliminary Amendment for filing in connection with the above-identified application.

- ☒ The Commissioner is hereby authorized to charge any fee, including any submitted herewith if the attached check(s) is in the wrong amount or otherwise improper or missing, that may be due in connection with this and the attached papers, or with this application during its entire pendency to or to credit any overpayment to Deposit Account No. 50-1213. A duplicate of this sheet is enclosed.

Respectfully submitted,  
HELLER EHRMAN WHITE & McAULIFFE LLP

By:

Stephanie Seidman  
Registration No. 33,779

Attorney Docket No. 24881-301D  
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#7

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PRELIMINARY AMENDMENT

Commissioner for Patents  
Arlington, VA 22202

Dear Sir:

Preliminary to examination of the above-captioned patent application, please  
amend the application as follows:

**IN THE SPECIFICATION:**

Please amend the specification as follows (a marked-up copy of the amended  
specification is attached to this Amendment):

**Please replace the paragraph beginning on page 3, line 26, through page 4, line  
8, with the following paragraph:**

Cells known to produce IL-1ra include monocytes, neutrophils, macrophages and  
fibroblasts. Cytokines known to upregulate IL-1Ra production include IL-13, IL-6, IL-4,  
IFN- $\gamma$ , GM-CSF and TGF- $\beta$ , the latter apparently by triggering IL-1 production which  
itself triggers IL-1Ra synthesis. The amino acid sequences of IL-1Ra from at least four  
species have been determined (human, rat, mouse and rabbit) and found to be at least  
75% homologous (Cominelli *et al.* (1994) *J. Biol. Chem.* 269:6963). IL-1Ra can also be  
synthesized as a strictly intracellular form whose production is the result of an  
alternative splicing of exon 1 (Butcher *et al.* (1994) *J. Immunol.* 153:701; Arend *et al.*  
(1993) *Adv. Immunol.* 54:167). IL-1Ra is released *in vivo* during experimentally-  
induced inflammation and as part of the natural course of many diseases. Administered  
experimentally, IL-1Ra has been demonstrated to block IL-1 activity *in vitro* and *in vivo*.

ai